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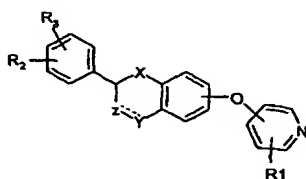
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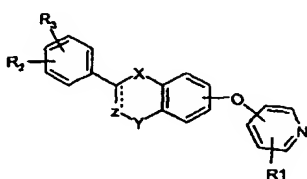
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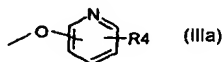
(54) Title: PYRIDINE DERIVATIVES USEFUL FOR INHIBITING SODIUM/CALCIUM EXCHANGE SYSTEM



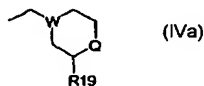
(I)



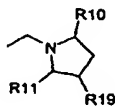
(II)



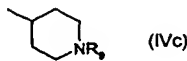
(IIIa)



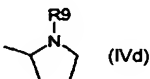
(IVa)



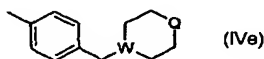
(IVb)



(IVc)



(IVd)



(IVe)

(57) Abstract: Therapeutically active compounds of formula (I) or (II) wherein X is -O-, -CH<sub>2</sub>- or -C(O)-; Z is -CHR<sub>12</sub>- or a valence bond; Y is -CH<sub>2</sub>-, -C(O)-, CH(OR<sub>13</sub>)-, -O-, -S-; provided that in case Z is a valence bond, Y is not C(O); the dashed line representing an optional double bond in which case Z is -CR<sub>12</sub>- and Y is -CH<sub>2</sub>-, -C(O)- or -CH(OR<sub>10</sub>)- (in formula II) or -CH- (in formula I); R<sub>2</sub> and R<sub>3</sub> are independently H, lower alkyl, lower alkoxy, -NO<sub>2</sub>, halogen, -CF<sub>3</sub>, -OH, benzyloxy or a group of formula (IIIa). R<sub>1</sub> is H, CN, halogen, -CONH<sub>2</sub>, -COOR<sub>15</sub>, CH<sub>2</sub>NR<sub>15</sub>R<sub>18</sub>, NHC(O)R<sub>5</sub>, NHCH<sub>2</sub>R<sub>5</sub>, NHR<sub>20</sub>, NR<sub>21</sub>R<sub>22</sub>, NHC(NH)NHCH<sub>3</sub> or, in case the compound is of formula (II) wherein the optional double bond exists or in case R<sub>2</sub> or R<sub>3</sub> is benzyloxy or a group of formula (IIIa) or in case the pyridine ring of formula (I) or (II) is attached to the oxygen atom in 3-, 4- or 5-position, R<sub>1</sub> can also be -NO<sub>2</sub> or NR<sub>16</sub>R<sub>17</sub>; R<sub>4</sub>

is H, -NO<sub>2</sub>, CN, halogen, -CONH<sub>2</sub>, -COOR<sub>15</sub>, -CH<sub>2</sub>NR<sub>15</sub>R<sub>18</sub>, -NR<sub>16</sub>R<sub>17</sub>, NHC(O)R<sub>5</sub> or -NHC(NH)NHCH<sub>3</sub>; R<sub>5</sub> is alkyl substituted with 1-3 substituents selected from the group consisting of halogen, amino and hydroxy, or carboxyalkyl, in which the alkyl portion is optionally substituted with 1-3 substituents selected from the group consisting of halogen, amino and hydroxyl, -CHR<sub>6</sub>NR<sub>8</sub> or one of the following groups: formula (IVa), (IVb), (IVc), (IVd), (IVe), and pharmaceutically acceptable salts and esters thereof. The compounds are potent inhibitors of Na<sup>+</sup>/Ca<sup>2+</sup> exchange mechanism.



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